ACTIVITIES OF JOINT USAGE/RESEARCH CENTER

JURC Cooperative Research Subjects 2012

(1 April 2012 ~ 31 March 2013)

STARTING-UP SUBJECTS (IN SPECIFIC FIELDS CHOSEN BY JURC)

Physics of THz Generation through the Interactions of Laser with Clusters

NAGASHIMA, Takeshi, Institute of Laser Engineering, Osaka University

Host in JURC HASHIDA, Masaki

Development of Beam Irradiation System with the Use of Electron Cooled Proton Beam and Its Application

SHIRAI, Toshiyuki, National Institute of Radiological Sciences **Host in JURC** NODA, Akira

Development of Methods to Measure Four Wave-mixing Process in Vacuum

HONMA, Kensuke, Graduate School of Science, Hiroshima University

Host in JURC SAKABE, Shuji

Development of Catalytic Enantioselective Reactions Induced by Hydrophobic Chiral Interface around Two-dimensionally Arrayed Gold Nanoparticles

MIKI, Kazushi, Polymer Materials Unit, National Institute for Materials Science

Host in JURC NAKAMURA, Masaharu

Optical and Electrical Properties of Mixed Organic Semiconductors NAITO, Hiroyoshi, The School of Engineering, Osaka Prefecture University

Host in JURC KANEMITSU, Yoshihiko

Development of Negative Thermal Expansion Material Based on a Perovskite BiNiO₂

AZUMA, Masaki, Materials and Structures Laboratory, Tokyo Institute of Technology

Host in JURC SHIMAKAWA, Yuichi

Synthesis of Sugar-Acetylenes by Iron-Catalyzed Cross-Coupling and Study on Their Stimulus-Response Luminescence

ORITA, Akihiro, Department of Applied Chemistry, Okayama University

Host in JURC NAKAMURA, Masaharu

Mechanistic Studies on the Iron Catalyze Carbon-carbon Bond Forming Reactions Based on the Solution-phase X-ray Absorption Spectroscopy

NAGASHIMA, Hideo, Institute for Materials Chemistry and Engineering, Kyushu University

Host in JURC NAKAMURA, Masaharu

Development of Ruthenium-Complex-Bound Amino Acids and Peptides and Their Application to Oxidative Degradation of Wooden Biomasses

WATANABE, Takashi, Research Institute for Sustainable Humanosphere, Kyoto University

Host in JURC NAKAMURA, Masaharu

Finding Frequent Similar Regions from Genome Sequences NAKAMURA, Atsuyoshi, Graduate School of Information Science and Technology, Hokkaido University

Host in JURC MAMITSUKA, Hiroshi

Efficient Search Algorithms for Structured Data in Bioinformatics TAKASU, Atsuhiro, National Institute of Informatics, Digital Content and Media Sciences Research Division

Host in JURC AKUTSU, Tatsuya

Probabilistic Methods for Analysis on Protein Interaction Networks

MARUYAMA, Osamu, Institute of Mathematics for Industry, Kyushu University

Host in JURC AKUTSU, Tatsuya

Development of Bioinformatic Method to Analyze MHC class II-binding Peptides

UDAKA, Keiko, Kochi Medical School **Host in JURC** MAMITSUKA, Hiroshi

Co-clustering of Biological Datasets for Personalized Medicine SHIGA, Motoki, Toyohashi University of Technology **Host in JURC** MAMITSUKA, Hiroshi

Development of the Evaluation Method for Potential Functionome Harbored in the Genome and Metagenome

TAKAMI, Hideto, Institute of Biogeoscience, Japan Agency for Marine-Earth Science and Technology

Host in JURC GOTO, Susumu

Development of Novel 3-Dimensional π -Extended Molecules Directed toward Electronic Materials

SUGA, Seiji, Graduate School of Natural Science and Technology, Okayama University

Host in JURC MURATA, Yasujiro

Evaluation of EO Effect of High Orientation Ba₂TiSi₂O₈ Thin Film by Prepared Sol-gel Method

IHARA, Rie, Department of Applied Physics, Tohoku University **Host in JURC** MASAI, Hirokazu

Investigation of Extraction Behaviors of Metal Ions in Liquidliquid Extraction Systems Using a Novel Multidentate Ligand MUKAI, Hiroshi, Faculty of Education, Kyoto University of Education

Host in JURC SOHRIN, Yoshiki

Role of Water in Motor Function of F1 Protein

IKEGUCHI, Mitsunori, Graduate School in Nanobioscience, Yokohama City University

Host in JURC MATUBAYASI, Nobuyuki

A Study on Intermolecular Atomic Contacts Involving Halogen and/or Chalcogen Atoms in Organic Crystals

TSUZUKI, Seiji, Nanosystem Research Institute, The National Institute of Advanced Industrial Science and Technology (AIST) **Host in JURC** SATO, Naoki

Electronic Properties of Thin Films of Porphyrazine Derivative Compounds Containing Thiadiazole Rings

AWAGA, Kunio, Research Center for Materials Science, Nagoya University

Host in JURC SATO, Naoki

Induction of Membrane Curvature by Arginine-rich Peptides SAKAMOTO, Kazutami, Faculty of Pharmacy, Chiba Institute of Science

Host in JURC FUTAKI, Shiroh

EXPANDING SUBJECTS (IN SPECIFIC FIELDS CHOSEN BY JURC)

X-Ray Structure Analysis of Reaction Mechanism of Catabolic Enzymes

OIKAWA, Tadao, Faculty of Chemistry, Materials and Bioengineering, Kansai University

Host in JURC HATA, Yasuo

Efficiency Improvement of Multi-dimensional Doppler Laser Cooling for Ultra-low Temperature Beam Generation OKAMOTO, Horomi, Graduate School of Advanced Sciences of Matter, Hiroshima University

Host in JURC NODA, Akira

X-Ray Structural Studies on Adaptation of Aspartate Rasemase to Environmental Temperatures

GOGAMI, Yoshitaka, Kansai University High Technology Research Center

Host in JURC HATA, Yasuo

Photocarrier Recombination Dynamics in Nanodomainstructured Ferroelectrics

FUNAKUBO, Hiroshi, Interdisciplinary Graduate School of Science and Engineering, Tokyo Institute of Technology Host in JURC KANEMITSU, Yoshihiko

Design and Synthesis of Metal Ligands Which Facilitate the Reductive Elimination Reaction and Their Application to the Synthesis of Cycloparaphenylenes

KORENAGA, Toshinobu, Graduate School of Natural Science and Technology, Okayama University **Host in JURC** YAMAGO, Shigeru

Efficient Construction of Polyfunctionalized Molecules by Cooperative Effect of Transition Metals and Heteroatoms SEGI, Masahito, School of Chemistry, College of Science and Engineering, Kanazawa University

Host in JURC TOSHIMITSU, Akio

Development of Methods for Discrimination and Syntheses of Chiral Molecules with Phosphoroselenoic Acid Derivatives Bearing a Binaphthyl Group

MURAI, Toshiaki, Faculty of Engineering, Gifu University Host in JURC TOSHIMITSU, Akio

Development of Reversible Carbon-Carbon Bond Formation Reactions on Dynamic Tetrametallic Reaction Sites OKAZAKI, Masaaki, Department of Frontier Materials Chemistry, Graduate School of Science and Technology, Hirosaki University Host in JURC OZAWA, Fumiyuki

Synthesis of Cyclic π -Conjugated Molecules and Their Properties SUZUKI, Toshiyasu, Institute for Molecular Science, National Institute of Natural Sciences

Host in JURC YAMAGO, Shigeru

Elucidation of the Mechanism of Encapsulation by Cycloparaphenylenes

HAINO, Takeharu, Graduate School of Science, Hiroshima University

Host in JURC YAMAGO, Shigeru

Cornea Regeneration by Small Molecule Fibronectin KOIZUMI, Noriko, Department of Biomedical Engineering, Faculty of Life and Medical Sciences, Doshisha University Host in JURC UESUGI, Motonari

Preparation of Organic-inorganic Hybrid Materials for Biosensor Application

FUJINO, Shigeru, Department of Chemical Engineering, Faculty of Engineering, Kyusyu University

Host in JURC YOKO, Toshinobu

Intracellular Gene Delivery Using Cell-penetrating Peptides KOGURE, Kentaro, Department of Biophysical Chemistry, Kyoto Pharmaceutical University

Host in JURC FUTAKI, Shiroh

Elucidation of Excitation Relaxation Dynamics and Application of Fullerenes Directly Connected to an Organic Dye OHKUBO, Kei, Department of Material and Life Science, Graduate School of Engineering, Osaka University

Host in JURC MURATA, Yasujiro

Synthesis, Structure and Electronic Properties of Carbazolophane-Polymers

TANI, Keita, Division of Natural Science, Osaka Kyoiku University Host in JURC TSUJII, Yoshinobu

Making of the Vertical Cross Section of Bioactive Trace Metals in the Japan Sea

NAKAGUCHI, Yuzuru, Faculty of Science and Engineering, Kinki University

Host in JURC SOHRIN, Yoshiki

Elucidation of Dewetting Process in Polymer Blend Thin Films as Studied by Grazing-Incidence Small Angle X-ray Scattering OGAWA, Hiroki, Japan Synchrotron Radiation Institute Host in JURC KANAYA, Toshiji

Investigation of Effect of Loop/Bridge Conformation Ratio on Elastic Properties of the BAB Triblock Copolymer/AB Ring Diblock Copolymer Having Middle Block with Type-A Dipoles Aligned Parallel to the Chain Contour

TAKAHASHI, Yoshiaki, Institute for Materials Chemistry and Engineering, Kyushu University

Host in JURC WATANABE, Hiroshi

Study of Segmental Dynamics in Multi-Component Polymer Systems in Miscible State

URAKAWA, Osamu, Graduate School of Science, Osaka University

Host in JURC MATSUMIYA, Yumi

Elucidation of Correlation between Structural Formation Process and Relaxation Process during Applying Shear Flow

MATSUBA, Go, Department of Polymer Science and Engineering, Faculty of Science and Engineering, Yamagata University Host in JURC KANAYA, Toshiji

Analysis of Molecular Adsorption of Water and Protein on a Stereo-Controlled Acrylamide Polymers

KATSUMOTO, Yukiteru, Graduate School of Science, Hiroshima University

Host in JURC HASEGAWA, Takeshi

STARTING-UP SUBJECTS (ON-DEMAND FROM RELATED COMMUNITIES)

Study on the Regulatory Mechanism of Plant Epidermal Cell Differentiation

TOMINAGA, Rumi, Interdisciplinary Research Organization, University of Miyazaki

Host in JURC AOYAMA, Takashi

Overexpression of Miraculin Using *Arabidopsis thaliana* INOUE, Hiroyasu, Faculty of Human Life and Environment, Nara Women's University

Host in JURC AOYAMA, Takashi

Small Molecules that Promote Differentiation into Pancreatic Beta Cells

KUME, Shouen, Institute of Molecular Embryology and Genetics, Kumamoto University

Host in JURC UESUGI, Motonari

Study for Tunnel Magnetoresistive Effect and Local Magnetism of Magnetic Tunnel Junctions Using Co₂MnSn Heusler Alloy Electrodes Prepared by Atomically-controlled Alternate Deposition TANAKA, Masaaki, Department of Engineering Physics, Electronics and Mechanics, Nagoya Institute of Technology **Host in JURC** ONO, Teruo

Synthesis and Functions of Multi-Bridged Naphthalene Oligomers

KURAMOCHI, Koji, Graduate School of Life and Environmental Sciences. Kyoto Prefectural University

Host in JURC KAWABATA, Takeo

High-energy Ion Generation through the Interactions of Laser with Nano Particles

NAKAI, Mitsuo, Institute of Laser Engineering, Osaka University **Host in JURC** SAKABE, Shuji

Identification and Functional Analysis of Bacterial Proteins Involved in Metal Metabolism

KURATA, Atsushi, Department of Applied Biological Chemistry, Faculty of Agriculture, Kinki University

Host in JURC KURIHARA, Tatsuo

Single-Electron Transistor Circuit Based on Nanoparticle MAJIMA, Yutaka, Materials & Structures Laboratory, Tokyo Institute of Technology

Host in JURC TERANISHI, Toshiharu

Programmable Architecture of Metal Complex-metal Nanoparticle Composites

TANAKA, Kentaro, Graduate School of Science, Nagoya University

Host in JURC TERANISHI, Toshiharu

Preparation, Structure, and Electronic Properties of Thin Films of D(donor)–A(acceptor)–D and A–D–A Type Compounds HAYASHI, Naoto, Graduate School of Science and Engineering for Research, University of Toyama

Host in JURC SATO, Naoki

EXPANDING SUBJECTS (ON-DEMAND FROM RELATED COMMUNITIES)

Efficient Synthesis of π -Conjugated Polymers via Direct Arylation KANBARA, Takaki, Graduate School of Pure and Applied Sciences, Tsukuba Research Center for Interdisciplinary Materials Science, University of Tsukuba

Host in JURC OZAWA, Fumiyuki

The Control of the Antiphase Boundary in Ferrimagnetic Spinel Ultrathin Films

NAGAHAMA, Taro, Laboratory of Advanced Materials Chemistry, Graduate School of Engineering, Hokkaido University **Host in JURC** ONO, Teruo

Optical Functionalities of Silicon Photonic Crystals TAKAHASHI, Yasushi, Reseach Organization for the 21st Century, Osaka Prefecture University

Host in JURC KANEMITSU, Yoshihiko

Search for Biologically Active Compounds from a Synthetic Library of Nitogen Heterocycles with Chiral Tetrasubstituted Carbon

ISHIBASHI, Masami, Graduate School of Pharmaceutical Sciences, Chiba University

Host in JURC KAWABATA, Takeo

Studies on the Transport and Metabolism of the Essential Trace Element Selenium in Mammals

MIHARA, Hisaaki, Department of Biotechnology, College of Life Sciences, Ritsumeikan University

Host in JURC KURIHARA, Tatsuo

Chemical Biology for Fine Analysis of Plant Hormone Signal Transduction Systems

HAYASHI, Ken-ichiro, Department of Biochemistry, Okayama University of Science

Host in JURC AOYAMA, Takashi

Synthesis and Properties of Well-Defined Poly(phenylene ethylene)s by Means of Organometalic Catalyst

KONISHI, Gen-ichi, Department of Organic and Polymeric Materials, Graduate School of Engineering, Tokyo Institute of Technology

Host in JURC NAKAJIMA, Yumiko

Dynamics of Spin-dependent Transport in Nanostructured

KOBAYASHI, Kensuke, School of Science, Osaka University **Host in JURC** ONO, Teruo

Electric and Magnetic Characteristics of New Ilmenite-type Iron

FUJII, Tatsuo, Faculty of Enginnering, Okayama University **Host in JURC** SAITO, Takashi

Mechanistic Study of Calpain Modulation by Bioinformatics ONO, Yasuko, Tokyo Metropolitan Institute of Medical Science **Host in JURC** MAMITSUKA, Hiroshi

Bulk Hetero Junction Photovoltaic Devices Composed of Novel Donar Polymer and Novel Fullerene Derivatives

IE, Yutaka, The Institute of Scientific and Industrial Research, Department of Soft Nanomaterials, Nanoscience and Nanotechnology Center, Osaka University

Host in JURC MURATA, Yasujiro

Chemical Biology of GGsTop, a Compound Inducing Antioxidative Stress Response of Human Skin Cells

KOJIMA, Akiko, Department of Food and Human Health Sciences, Graduate School of Human Life Sciences, Osaka City University

Host in JURC HIRATAKE, Jun

SUBJECTS FOCUSING OF JOINT USAGE OF JURC/ICR FACILITIES

Nano-electron Spectroscopic Study of Helium Bubbles in Si MIYAMOTO, Mitsutaka, Interdisciplinary Faculty of Science and Engineering, Shimane University

Host in JURC KURATA, Hiroki

Systematic Syntheses and Properties of Polycyclic Aromatics Containing a Phosphorus Atom

OKUMA, Kentaro, Department of Chemistry, Faculty of Science, Fukuoka University

Host in JURC SASAMORI, Takahiro

Physicochemical Properties of Novel Aromatic Compounds with Heteroatoms

SAITO, Masaichi, Department of Chemistry, Graduate School of Science and Engineering, Saitama University

Host in JURC TOKITOH, Norihiro

Synthesis and Structure of Transition Metal Complexes with New Tripodal Tetradentate Ligand

UNNO, Masafumi, Graduate School of Engineering, Gunma University

Host in JURC TOKITOH, Norihiro

Synthesis and Structure of Main Group Element-Protected Metal Nanoparticles

FUJIHARA, Hisashi, Department of Chemistry, Fuculty of Science and Engineering, Kinki University

Host in JURC TOKITOH, Norihiro

Synthesis and Structure of Color-tunable Organotelluronium Salts

MINOURA, Mao, School of Science, Kitasato University **Host in JURC** TOKITOH, Norihiro

Structural Features of Aromatic-rings-accumulated Molecules in Crystal

OKAMOTO, Akiko, Department of Organic and Polymer Materials Chemistry, Tokyo University of Agriculture & Technology **Host in JURC** NAKAMURA, Masaharu

Solid-State NMR Analysis of Molecular Orientation in Organic LED Materials

ADACHI, Chihaya, OPERA, Kyushu University **Host in JURC** KAJI, Hironori

NMR Characterization of Carbon Nitride TAKARABE, Kenichi, Department of Applied Science, Okayama University of Science Host in JURC KAJI, Hironori

Solid-state-NMR Study on Luminescence and Structure Characteristics of Organic Molecules

TOKUDOME, Yasuaki, Graduate School of Engineering, Osaka Prefecture University

Host in JURC KAJI, Hironori

SUBJECTS ENCOURAGING JOINT PROGRAM

Elucidation of Electronic States, Local Structures, and Properties in Transition Metal Oxides Using Synchrotron Radiated X-rays MIZUMAKI, Masaichiro, Japan Synchrotron Radiation Research Institute, SPring-8

Host in JURC ICHIKAWA, Noriya

The 8th International Workshop for East Asian Young Rheologists

TAKAHASHI, Yoshiaki, Institute for Materials Chemistry and Engineering, Kyushu University

Host in JURC WATANABE, Hiroshi

JURC Publications

(until 31 May 2012)

Dielectric and Viscoelastic Investigation of Segmental Dynamics of Polystyrene above Glass Transition Temperature: Cooperative Sequence Length and Relaxation Mode Distribution Matsumiya, Y.; Uno, A.; Watanabe, H., *Macromolecules*, 44, 4355-4365 (2011).

Abstract

Atactic polystyrene (PS) has the type-B dipole perpendicular to the chain backbone so that its local, segmental motion activates the dielectric relaxation. For monodisperse oligostyrene (OS) and PS samples of various molecular weights M, details of this motion were examined at temperatures T well above T_{g} through comparison of the complex modulus, $G^* = G' + iG''$, and the complex dielectric permittivity, $\varepsilon^* = \varepsilon' - i\varepsilon''$, measured as functions of the angular frequency ω . For the OS samples, $G^*(\omega)$ and $\varepsilon^*(\omega)$ fully relaxed through the segmental dynamics thereby exhibiting respective terminal relaxation tails (low-frequency tails), $G'(\omega) \propto \omega^2$, $G''(\omega) \propto \omega$, $\Delta \varepsilon'(\omega) \equiv \varepsilon'(0) - \varepsilon'(\omega) \propto \omega^2$, and $\varepsilon''(\omega) \propto \omega$, at ω below the segmental relaxation frequency ω_s . For the PS samples, $G^*(\omega)$ relaxed partly through the segmental dynamics and then exhibited the polymeric full relaxation characterized by the Rouse-like behavior followed by the terminal flow behavior (with/without intermediate entanglement plateau depending on M). In contrast, $\varepsilon^*(\omega)$ of the PS samples still relaxed completely through the segmental dynamics. For respective samples, the $G^*(\omega)$ and $\varepsilon^*(\omega)$ data in the segmental relaxation zone exhibited very similar relaxation mode distribution and had the same time—temperature shift factor. Nevertheless, a ratio of the dielectrically and viscoelastically detected segmental relaxation times, $r(M) = \omega_{s,G}/\omega_{s,\varepsilon}$, and the dielectric relaxation intensity, $\Delta \varepsilon(M)$, decreased with increasing M up to $M^* = 2 \times 10^3$ and then became insensitive to M on a further increase of M. The viscoelastic segmental relaxation reflects the cooperative torsion of the repeating units along the molecular backbone (as noted from rheo-optical data), while the dielectric segmental relaxation detects reorientational motion of those units affected by both intra- and intermolecular cooperativity (as noted from the basic dielectric expression). The observed decreases of r(M) and $\Delta \varepsilon(M)$ suggested that the dimension ξ_m of the whole OS molecule (over which the cooperative torsion occurs) is smaller than the length scale ξ_c for the intermolecular cooperative motion and that ξ_m approaches ξ_c on an increase of M up to M^* . Consequently, the high-M PS molecules having $\xi_{\rm m} > \xi_{\rm c}$ exhibited the M-insensitive r(M) and $\Delta \varepsilon(M)$. Thus, the M value for the crossover between these two regimes, $M^* = 2 \times 10^3$, can be taken as the molecular weight of the cooperative sequence along the PS backbone. Furthermore, the quantitative similarity of the viscoelastic and dielectric mode distributions suggests that the cooperative torsion of the repeating units along the molecular backbone is governed by the cross-correlation of the units belonging to different molecules.

Directional Terahertz Emission from Air Plasma Generated by Linearly Polarized Intense Femtosecond Laser Pulses

Jahangiri, F.; Hashida, M.; Tokita, S.; Nagashima, T.; Ohtani, K.; Hangyo, M.; Sakabe, S., *Applied Physics Express*, **5**, [026201-1]-[026201-3] (2012).

Abstract

Terahertz (THz) radiation from air plasma produced by linearly polarized intense femtosecond laser pulses was investigated. The laser energy dependence, directionality, and polarization properties of THz waves, measured in the present experiment, differed from those in previous reports and can be explained by parametric decay of laser light to R-waves in the presence of a spontaneous magnetic field.

Colossal Negative Thermal Expansion in BiNiO₃ Induced by Intermetallic Charge Transfer

Azuma, M.; Chen, W. T.; Seki, H.; Czapski, M.; Olga, S.; Oka, K.; Mizumaki, M.; Watanuki, T.; Ishimatsu, N.; Kawamura, N.; Ishiwata, S.; Tucker, M. G.; Shimakawa, Y.; Attfield, J. P., *Nature Communications*, [DOI: 10.1038/ncomms1361] (2011).

Abstract

The unusual property of negative thermal expansion is of fundamental interest and may be used to fabricate composites with zero or other controlled thermal expansion values. Here we report that colossal negative thermal expansion (defined as linear expansion $< -10^{-4} \text{ K}^{-1}$ over a temperature range $\sim 100 \text{ K}$) is accessible in perovskite oxides showing charge-transfer transitions. BiNiO₃ shows a 2.6% volume reduction under pressure due to a Bi/Ni charge transfer that is shifted to ambient pressure through lanthanum substitution for Bi. Changing proportions of coexisting low- and high-temperature phases leads to smooth volume shrinkage on heating. The crystallographic linear expansion coefficient for $Bi_{0.95}La_{0.05}NiO_3$ is -137×10^{-6} K⁻¹ and a value of -82×10^{-6} K⁻¹ is observed between 320 and 380 K from a dilatometric measurement on a ceramic pellet. Colossal negative thermal expansion materials operating at ambient conditions may also be accessible through metal-insulator transitions driven by other phenomena such as ferroelectric orders.

Preparation of Co₂FeSn Heusler Alloy Films and Magnetoresistance of Fe/MgO/Co₂FeSn Magnetic Tunnel Junctions

Tanaka, M. A.; Ishikawa, Y.; Wada, Y.; Hori, S.; Murata, A.; Horii, S.; Yamanishi, Y.; Mibu, K.; Kondou, K.; Ono, T.; Kasai, S., *Journal of Applied Physics*, **111**, [053902-1]-[053902-4] (2012).

Abstract

To obtain magnetic tunnel junctions (MTJs) composed of non-equilibrium alloy, Co₂FeSn films were prepared by atomically controlled alternate deposition at various substrate temperatures. X-ray diffraction patterns and Mössbauer spectra clarify that Co₂FeSn films in the Heusler alloy phase can be realized by growing at a substrate temperature of 250 °C or below. Phase separation into cubic CoSn, hexagonal CoSn and cubic CoFe phases occurs in films grown at substrate temperatures 300 °C or greater. Fe/MgO/Co₂FeSn MTJs were prepared with the Co₂FeSn layer grown at various substrate temperatures. The MTJs with the ferromagnetic Co₂FeSn layer grown at a substrate temperature of 250 °C showed tunnel magnetoresistance ratios of 72.2% and 43.5% at 2 K and 300 K, respectively.

Nanoparticles for *ex vivo* siRNA Delivery to Dendritic Cells for Cancer Vaccines: Programmed Endosomal Escape and Dissociation

Akita, H.; Kogure, K.; Moriguchi, R.; Nakamura, Y.; Higashi, T.; Nakamura, T.; Serada, S.; Fujimoto, M.; Naka, T.; Futaki, S.; Harashima, H., *Journal of Controlled Release*, **143**, 311-317 (2010).

Abstract

We previously developed octaarginine (R8)-modified lipid envelope-type nanoparticles for siRNA delivery (R8-MEND). Herein, we report on their *ex vivo* siRNA delivery to primary mouse bone marrow-derived dendritic cells (BMDCs) for poten-

tial use as a cancer vaccine. Quantitative imaging analysis of the intracellular trafficking of siRNA revealed that the dissociation process, as well as the rate of endosomal escape limits the siRNA efficiency of the prototype R8-MEND, prepared by the hydration method (R8-MEND_{hydo}). Successful endosomal escape was achieved by using a pH-dependent fusogenic peptide (GALA) modified on a lipid mixture that was optimized for endosomal fusion. Furthermore, a modified protocol for the preparation of nanoparticles, mixing the siRNA/STR-R8 complex and small unilamellar vesicles (R8/GALA-MEND_{SUV}), results in a more homogenous, smaller particle size, and results in a more efficient intracellular dissociation. Gene knockdown of the suppressor of cytokine signaling 1 (SOCS1), a negative feedback regulator of the immune response in BMDCs resulted in an enhanced phosphorylation of STAT1, and the production of proinflammatory cytokines. Moreover, SOCS1-silenced BMDCs were more potent in suppressing tumor growth. Collectively, these results show that siRNA loaded in R8/GALA-MEND_{SUV} efficiently suppresses endogenous gene expression and consequently enhances dendritic cell-based vaccine potency in vivo.